AMENDMENTS TO THE CLAIMS

This listing of claims will replace all prior versions, and listings, of claims in the application:

1. (original) Use of a compound of the formula (I), or a pharmaceutically acceptable salt thereof in the manufacture of a medicament for use in the treatment or prevention of a condition involving sodium ion flux through a sensory neurone specific channel of a sensory neurone

$$(R_1)_n \xrightarrow{X} R_2$$

$$(R_1)_n \xrightarrow{(CH_2)_m} (I)$$

wherein:

- X is -N- or -CH-;
- n is from 0 to 3;
- each R₁ is the same or different and is a hydroxy, amino, halogen, C₁-C₆ alkyl, C₁-C₆ haloalkyl, C₁-C₆ alkoxy, C₂-C₆ alkenyloxy, C₂-C₆ alkynyloxy, C₁-C₆ haloalkoxy, C₁-C₆ alkylthio, C₁-C₆ haloalkylthio, (C₁C₆ alkyl) amino or di (C₁-C₆ alkyl) amino group;
- p is 0 or 1;
- R_1^{-1} is cyano, -NR₁-CO-(C₁-C₄ alkyl), -NR₁-S(O)₂-(C₁-C₄ alkyl), -CO₂H, -S(O)₂OH, -CO₂-(C₁-C₄ alkyl), -O-S(O)₂-(C₁-C₄ alkyl) or -N[S(O)₂-(C₁-C₄ alkyl)]₂, wherein R₁ is hydrogen or a C₁-C₄ alkyl group;
- m is 1, 2 or 3; and
- R₂ is either

- (a) -L-A, wherein L is a direct bond or a C_1 - C_6 alkyl, C_2 - C_6 alkenyl or C_2 - C_6 alkynyl moiety and A is C_6 - C_{10} aryl, C_3 - C_6 carbocyclyl, a 5- to 10-membered heteroaryl group or a 5- to 10- membered heterocyclic group,
- (b) -L-CR(A)₂ or -L-CH=C(A)₂ wherein R is hydrogen or C₁-C₄ alkyl, L is as defined above and each A is the same or different and is as defined above,
- (c) $-L^1$ -Het-A¹, wherein Het is -O-, -S- or $-NR^1$ -, A¹ is -L-A, -L-CR(A)₂ or -L-CH=C (A)₂, R¹ is H or -L-A, L¹ is a C₁-C₆ alkyl, C₂-C₆ alkenyl or C₂-C₆ alkynyl moiety, L is as defined above, R is as defined above and each A is the same or different and is as defined above,
- -L-CO-NR₃R₄ or -L-CS-NR₃R₄, wherein L is as defined above and either (i) R₃ and R₄, together with the N atom to which they are attached, form a 5- to 10-membered heteroaryl or heterocyclyl group or (ii) R₃ represents -L-H or A¹ wherein L and A¹ are as defined above, and R₄ represents -L¹-H,-L¹-CO-A¹,-L¹-S (O)-A¹, -L¹-S(O)₂-A¹, -L¹-Het-A¹, -NR-CO-N(A)₂, -N(A)₂,-A-Het-A, -A¹, -L-CR(LA)₂ or -L-CH=C(LA)₂ wherein each L is the same or different, each A is the same or different, and L¹, L, R, A and A¹ are as defined above,
- (e) -CO-L-NR₃R₄ or -CS-L-NR₃R₄ wherein L, R₃ and R₄ are as defined above, (f) -CO-A¹ or -CS-A¹ wherein A¹ is as defined above,
- (g) $-L^1$ -O-N=C(A)₂ or -CO-L¹-O-N=C(A)₂ wherein L¹ is as defined above and each A is the same or different and is as defined above, or
- (h) -L¹-NR-CO-NR₃R₄ or -L¹-NR-CS-NR₃R₄, wherein L¹, R, R₃ and R₄ are as defined above,

wherein

- said aryl, carbocyclyl, heteroaryl and heterocyclyl groups are optionally fused to one or two cyclic moieties selected from phenyl rings and 5- to 6-membered heterocyclyl and heteroaryl groups, and
- said aryl, heteroaryl, carbocyclyl and heterocyclyl groups are unsubstituted or are substituted by 1, 2 or 3 substituents which are the same or different and are selected from C_1 - C_4 alkyl, C_1 - C_4 haloalkyl, halogen, hydroxy, amino, $(C_1$ - C_4 alkyl)amino, $di(C_1$ - C_4 alkyl)amino, C_1 - C_4 alkoxy, C_1 - C_4 haloalkoxy, C_1 - C_4 alkylthio, C_1 - C_4 haloalkylthio, -NH-CO- $(C_1$ - C_4 alkyl), -CO- $(C_1$ - C_4) alkyl, -CO₂- $(C_1$ - C_4 alkyl), 5-or 6- membered heteroaryl, phenyl and -CHPh₂ substituents, the phenyl and heteroaryl moieties in said substituents being unsubstituted or substituted by 1 or 2 further substituents selected from halogen atoms, C_1 - C_2 alkyl groups, C_1 - C_2 alkoxy groups and -NH-CO- $(C_1$ - C_2 alkyl) groups,

provided that (a) when R₂ is -L-A, A is other than a benzimidazolyl group, and (b) when R₂ is -CO-A¹ or -CS-A¹, A is other than a pyrazolopyrimidinyl or pyrazolyl group.

- 2. (original) Use according to claim 1, wherein:
- X is -N- or -CH-;
- n is from 0 to 3;
- p is 0;
- each R₁ is the same or different and is a hydroxy, amino, halogen, C₁-C₆ alkyl, C₁-C₆ haloalkyl, C₁-C₆ alkoxy, C₁-C₆ haloalkoxy, C₁-C₆ alkylhio, C₁-C₆ haloalkylthio, (C₁C₆ alkyl)amino or di(C₁-C₆ alkyl)amino group;

- m is 1, 2 or 3; and
- R₂ is either
- (a) -L-A, wherein L is a direct bond or a C₁-C₆ alkyl, C₂-C₆ alkenyl or C₂-C₆ alkynyl moiety and A is C₆-C₁₀ aryl, C₃-C₆ carbocyclyl, a 5- to 10-membered heteroaryl group or a 5- to 10-membered heterocyclic group,
- (b) -L-CR(A)₂ or-L-CH=C(A)₂ wherein R is hydrogen or C₁-C₄ alkyl, L is as defined above and each A is the same or different and is as defined above,
- (c) $-L^1$ -Het-A¹, wherein Het is -O-, -S- or $-NR^1$ -, A¹ is -L-A, -L-CR(A)₂ or -L-CH=C (A)₂, R¹ is H or -L-A, L1 is a C₁-C₆ alkyl, C₂-C₆ alkenyl or C₂-C₆ alkynyl moiety, L is as defined above, R is as defined above and each A is the same or different and is as defined above,
- (d) -L-CO-NR₃R₄ or-L-CS-NR₃R₄, wherein L is as defined above and either (i) R₃ and R₄, together with the N atom to which they are attached, form a 5- to 10-membered heteroaryl or heterocyclyl group or (ii) R₃ represents -L-H or A¹ wherein L and A¹ are as defined above, and R₄ represents -L¹-H,-L¹-CO-A, A¹,-L-CR (LA)₂ or -L-CH=C (LA)₂ wherein each L is the same or different, each A is the same or different, and L¹, L, R, A and A¹ are as defined above,
- (e) -CO-L-NR₃R₄ or -CS-L-NR₃R₄ wherein L, R₃ and R₄ are as defined above, (f) -CO-A¹ or -CS-A¹ wherein A¹ is as defined above, or
- (g) -L-O-N=C(A)₂ or -CO-L-O-N=C(A)₂ wherein L is as defined above and each A is the same or different and is as defined above, wherein

- said aryl, carbocyclyl, heteroaryl and heterocyclyl groups are optionally fused to one or two cyclic moieties selected from phenyl rings and 5- to 6- membered heterocyclyl and heteroaryl groups, and
- said aryl, heteroaryl, carbocyclyl and heterocyclyl groups are unsubstituted or are substituted by 1, 2 or 3 substituents which are the same or different and are selected from C₁-C₄ alkyl, C₁-C₄ haloalkyl, halogen, hydroxy, C₁-C₄ alkoxy, C₁-C₄ haloalkylthio, phenyl and-CHPh2 substituents, the phenyl moieties in said substituents being unsubstituted or substituted by 1 or 2 halogen atoms,

provided that (a) when R_2 is -L-A, A is other than a benzimidazolyl group and (b) when R_2 is -CO-A¹ or -CS-A¹, A is other than a pyrazolopyrimidinyl or pyrazolyl group.

3. (currently amended) Use according to claim 1-or-2, wherein the aryl, heteroaryl, heterocyclyl and carbocyclyl groups and moieties in the substituents R₁, R₂, R₃ and R₄ are unsubstituted or substituted by 1, 2 or 3 substituents which are the same or different and are selected from halogen, C₁-C₄ alkyl, hydroxy, amino, (C₁-C₄ alkyl) amino, C₁-C₄ alkoxy, C₁-C₄ haloalkyl, C₁-C₄ haloalkoxy, C₁-C₄ alkylthio, C₁-C₄ haloalkylthio, -NH-CO-(C₁-C₂ alkyl), -CO-(C₁-C₂ alkyl), -CO₂-(C₁-C₂ alkyl), 5-membered heteroaryl, phenyl and-CHPh₂ substituents, the phenyl and heteroaryl moieties in said substituents being unsubstituted or substituted by one or two further substituents selected from halogen atom, C₁-C₂ alkyl groups, C₁-C₂ alkoxy groups and -NH-CO-(C₁-C₂ alkyl) groups.

- 4. (currently amended) Use according to any one of the preceding claims claim 1, wherein each R₁ is the same or different and is a hydroxy, amino, halogen, C₁-C₄ alkyl, C₁-C₄ haloalkyl, C₁-C₄ alkoxy, C₂-C₄ alkenyloxy, C₁-C₄ haloalkoxy, C₁-C₄ alkylthio or C₁-C₄ haloalkylthio group.
- 5. (currently amended) Use according to any one of the preceding claims claim 1, wherein each L moiety in the R_2 substituent is the same or different and represents a direct bond or a C_1 - C_4 alkyl moiety and/or each L^1 moiety in the R_2 substituent is the same or different and represents a C_1 - C_4 alkyl moiety.
- 6. (currently amended) Use according to <u>any one of the preceding claims claim 1</u>, wherein each A moiety in the R₂ substituent is the same or different and represents a C₆-C₁₀ aryl, C₃-C₆ cycloalkyl, 5-or 6-membered heterocyclyl or 5-or 6-membered heteroaryl group, which group is (a) unsubstituted or substituted by 1, 2 or 3 substituents selected from C₁-C₄ alkyl, C₁-C₄ haloalkyl, halogen, hydroxy, amino, (C₁-C₄ alkyl)amino, di (C₁-C₄alkyl)amino, C₁-C₄ alkoxy, C₁-C₄ haloalkoxy, C₁-C₄ alkylthio, C₁-C₄ haloalkylthio, -NH-CO-(C₁-C₂ aLkyl), phenyl and halophenyl substituents and (b) optionally fused to one or two cyclic moieties selected from phenyl rings and 5- to 6-membered heterocyclyl or heteroaryl groups.

- 7. (currently amended) Use according to any one of the preceding claims claim 1, wherein each R substituent in each -CR(A)₂ moiety is the same or different and is hydrogen or methyl.
- 8. (currently amended) Use according to any one of the preceding claims claim 1, wherein each Het moiety in the R₂ substituent is -O-, -S- or -NR- wherein R is hydrogen, C₁-C₄ alkyl, phenyl or -(C₁-C₄ alkyl)-phenyl.
- 9. (currently amended) Use according to any one of the preceding claims claim 1, wherein, when R₃ and R₄, together with the nitrogen atom to which they are attached, form a heterocycle, they form a 5- to 7- membered heterocyclyl group.
- 10. (original) Use according to claim 9, wherein, when R₃ and R₄, together with the nitrogen atom to which they are attached, form a heterocycle, they form a morpholino, thiomorpholino, S-oxo-thiomorpholino, S,S-dioxo-thiomorpholino, pyrrolidinyl, piperazinyl or homopiperidinyl ring which is (a) optionally fused to one or two cyclic moieties selected from phenyl rings and 5- to 6- membered heteroaryl rings, and (b) unsubstituted or substituted by 1 or 2 substituents selected from C₁-C₄ alkyl, C₁-C₄ haloalkyl, C₁-C₄ alkoxy, C₁-C₄ alkylthio, halogen, phenyl, -CHPh₂, -CO-(C₁-C₂ alkyl), -CO₂-(C₁-C₂ alkyl) and 5-to 6-membered heteroaryl substituents, the phenyl and heteroaryl moieties in said substituents being unsubstituted or substituted by 1 or 2

further substituents selected from halogen atoms, C_1 - C_2 alkyl groups, C_1 - C_2 alkoxy groups and -NH-CO (C_1 - C_2 alkyl) groups.

- 12. (currently amended) Use according to any one of the preceding claims claim 1, wherein, when R₂ is defined according to option (a), A is monocyclic.
- 13. (currently amended) Use according to any one of the preceding claims claim 1, wherein, when R_2 is defined according to option (f), A is a said C_6 - C_{10} aryl group.

- 14. (currently amended) Use according to any one of the preceding claims claim 1, wherein
- X is -N- or -CH-;
- n is 0 or 1;
- each R_1 is the same or different and is C_1 - C_2 alkyl, hydroxy or C_1 - C_2 alkoxy;
- p is 0 or 1;
- R₁¹ is cyano, -NH-CO-CH₃, -NH-S(O)2-CH₃, -O-S(O)₂-CH₃, -N[SO₂-CH₃]₂ or -S(O)₂-OH;
- m is 1, 2 or 3; and
- R₂ is either
- -L-A wherein L represents a direct bond or a C₁-C₄ alkyl moiety, for example a methyl, ethyl or propyl moiety, and A is a phenyl, thienyl, triazolyl, pyridyl, fluorenyl, thiazolyl, tetrahydroisoquinolinyl, 9H-carbazolyl, indolinyl, 9H- xanthenyl or benzimidazolyl group, which group is unsubstituted or substituted by one or two substituents selected from halogen, C₁-C₂ alkyl, hydroxy, amino, C₁-C₂ alkoxy, C₁-C₂ haloalkyl, C₁-C₂haloalkoxy, C₁-C₂ haloalkylthio, -NH-CO-CH₃ and phenyl substituents,
- (b) -L-CR(A) 2 or -L-CH=C(A)₂ wherein R is hydrogen or methyl, L is as defined above and each A is the same or different and is as defined above,
- (c) -L¹-Het-A¹ wherein Het is -O-or –NR¹-wherein R¹ is hydrogen, C₁-C₄ alkyl or benzyl, A¹ is -L-A, -L-CR(A)₂ or -L-CH=C(A)₂, L¹ is a C₁-C₄ alkyl moiety, for example a methyl, ethyl or propyl moiety, L is as defined above, R is as defined above and each A is the same or different and is as defined above,

-L-CO-NR₃R₄ wherein L is as defined above and either (i)R₃ and R₄, together (d) with the nitrogen atom to which they are attached, form a morpholino, thiomorpholino, S-oxo-thiomorpholino, S.S-dioxo-thiomorpholino, pyrrolidinyl, piperazinyl or homopiperidinyl ring which is (a) optionally fused to one or two cyclic moieties selected from phenyl rings and 5-to 6- membered heteroaryl rings, and (b) unsubstituted or substituted by one or two substituents selected from C₁-C₄ alkyl, C₁-C₄ haloalkyl, C₁-C₄ alkoxy, C₁-C₄ alkylthio, halogen, phenyl, -CHPh₂, -CO-(C₁-C₂ alkyl), -CO₂-(C₁-C₂ alkyl) and 5-to 6-membered heteroaryl substituents, the phenyl and heteroaryl moieties in said substituents being unsubstituted or substituted by one or two further substituents selected from halogen atoms, C₁-C₂ alkyl groups, C₁-C₂ alkoxy groups and -NH-CO-(C₁-C₂ alkyl) groups, or (ii) R₃ represents hydrogen, C₁-C₄ alkyl or an unsubstituted benzyl, phenyl, hydroxyphenyl or -(Ci-C2 alkyl) -CHPh₂ group and R₄ represents-C₁-C₄ alkyl, fluorenyl, phenyl, pyridyl, (C₁-C₄ alkyl)-phenyl, -(C₁-C₄ alkyl)-(5- to 6-membered heteroaryl), -(CH₂)_m-(9H-carbazolyl),-(CH₂)_m-indolinyl,-(CH₂)_m-(9H-xanthenyl), -(CH₂)_m-O-CHA¹¹ A¹¹¹, -(CH₂)_m-S-CHA¹¹ A¹¹¹, -(CH₂)_m-S(O)-CHA¹¹ A¹¹¹, -(CH₂)_m-S(O)₂-CHA¹¹ A¹¹¹, -NH-CO-N(phenyl)₂, -N(phenyl)₂ or -A¹¹-O-A¹¹¹, -(CH₂)_m-CHA¹¹ A¹¹¹, - $CH[(CH_2)_nPh]_2$ or $-(CH_2)_n-CO-R$ where m is 0, 1, 2 or 3, A^{11} and A^{111} are the same or different and each represent phenyl or a 5- or 6- membered heteroaryl group, n is 0, 1 or 2, p is 1,2 or 3 and R is 5- or 6- membered heterocyclic group fused to a phenyl ring, for example a- tetrahydroisoquinoline group, the cyclic moieties in said R₄ groups being unsubstituted or substituted by a halogen atom, C₁-C₂ alkyl, hydroxy, amino or C₁-C₂ alkoxy group,

- (e) -CO-L-NR₃R₄ or-CS-L-NR₃R₄ wherein L, R₃ and R₄ are as defined above, (f) -CO-A¹ or -CS-A¹ where A¹ is as defined above,
- (g) -CO-L¹-O-N=C(A)₂ wherein L¹ is as defined above and each A is the same or different and is as defined above; or
- (h) -L¹-NR-CO-NR₃R₄ or -L¹-NR-CS-NR₃R₄ wherein L¹, R, R3 and R4 are as defined above,

provided that when R₂ is -L-A, A is monocyclic.

- 15. (currently amended) Use according to any one of the preceding claims claim 1, wherein said condition is chronic or acute pain, a bowel disorder, a bladder dysfunction, tinnitus or a demyelinating disease.
- 16. (currently amended) A compound of the formula (I), as defined in any one of claims 1 to 14 claim 1, or a pharmaceutically acceptable salt thereof.
- 17. (currently amended) A pharmaceutical composition comprising a compound of the formula (I), as defined in any one of claims 1 to 14 claim 1, or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier or diluent.
- 18. (original) A composition according to claim 17 which is a capsule or tablet comprising from 10 to 500 mg of a compound of the formula (I), as defined in any one of claims 1 to 14, or a pharmaceutically acceptable salt thereof.

- 19. (original) An inhalation device comprising a pharmaceutical composition according to claim 18.
- 20. (original) An inhalation device according to claim 19 which is a nebulizer.
- 21. (currently amended) A compound according to any one of claims 1 to 14 claim 1, or a pharmaceutically acceptable salt thereof, for use in the treatment of the human or animal body.
- 22. (currently amended) A method of treating a patient suffering from or susceptible to a condition as defined in claim 1 or 15, which method comprises administering to said patient an effective amount of a compound of formula (I), as defined in any of claims 1 to 14, or a pharmaceutically acceptable salt thereof.